

Ebola survivors can pass on the virus: we're trying to understand what role sex plays

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The <u>current Ebola disease outbreak</u> in the Democratic Republic of the Congo (DRC) is the second largest in history. It has been raging for over a year with no indication that it is nearing an end. Many of the issues that prevent effective containment of Ebola disease outbreaks <u>have been discussed</u>. These include <u>porous borders</u>, <u>community mistrust</u> of health workers, and the spread of <u>misinformation</u>.



A health worker spreading disinfectant at a health checkpoint in Goma, DRC. Patricia Martinez/EPA-EFE

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But one area that requires further study is Ebola virus' persistence in the reproductive systems of otherwise healthy survivors. Understanding how Ebola disease outbreaks are exacerbated by sexual transmission is important to stop new chains of transmission and to prevent the disease from spreading to new geographic regions.

Multiple reports strongly suggest that <u>re-emergence</u> of Ebola disease is linked to <u>persistent infections</u> and <u>sexual</u> <u>transmission</u> from survivors. What this means is that people who have survived infection by Ebola virus – and who no longer show any symptoms of the disease – can continue to carry the virus and pass on the infection months after they have recovered.

Our research group recently <u>reviewed</u> the current understanding of rates and mechanisms of Ebola virus persistence in male disease survivors. We are using this information to fill in gaps about how persistence relates to sexual transmission of the virus.

In addition, we are combining molecular investigations with insights from survivors to better equip future outbreak response efforts. We also hope the work will help address long-term health issues faced by Ebola disease survivors.

The role of the reproductive system

Nearly everything known about Ebola virus persistence in the reproductive system has resulted from testing semen of West African Ebola disease survivors.

It has been established that the Ebola virus can be detected in semen long after it is cleared from other organs and tissues. For example, a <u>recent study</u> showed that the virus could be detected in <u>50% of male survivors 115 days</u> after recovery and may linger for up to three years.

The virus has also been detected in semen at concentrations greatly exceeding that found in the blood at peak infection. The <u>data</u> suggests that this occurs in the absence of any symptoms of disease. In addition, <u>sexual transmission</u> of Ebola virus from <u>male survivors</u> has been reported up to 18 months following their recovery.

These observations are important to public health. Firstly, long-term persistence of Ebola virus could lead to the initiation of new chains of disease transmission long after outbreaks have ended. Secondly, the presence of high concentrations of virus in the semen in the absence of disease symptoms means that male survivors could carry and transmit the virus without knowing they're still infected.

But the fact that only semen is being used in these studies presents a problem. This means that the perseverance of the virus is only being traced in males. This means that little is known about persistence of the virus in women. This urgently needs attention because Ebola virus has been detected in <u>female survivors</u> up to 15 months <u>following recovery</u>, and there are <u>recorded cases of female to male</u> transmission.

Unanswered questions

There is a strong need for greater understanding of Ebola virus infections, including:

- When Ebola virus is transmitted to reproductive tissues during illness.
- How Ebola virus persists over extended periods in reproductive tissues.
- How long persistent virus remains infectious.
- How often sexual transmission occurs.

In a bid to answer some of these questions – particularly the molecular intricacies of Ebola virus persistence within the reproductive system – we used laboratory models of male and female reproductive systems. This allowed us to:

- Study the dynamics of Ebola virus transmission to the reproductive system.
- Identify the particular types of cells that are infected by Ebola virus in the reproductive system.
- Determine how local immune responses in the infected cells help facilitate Ebola virus persistence.
- Characterise damage to the reproductive system, including altered sperm health.

We have also performed Ebola virus infections in mice to better understand how our molecular data reflects the complex pathophysiology of Ebola virus disease and persistence in humans. Mouse models have been used in <u>numerous</u>

investigations of Ebola virus infection.

By infecting mice with Ebola virus, we can determine whether the male reproductive system is damaged by persistent infection. We can also determine if there is an effect on fertility, and how often persistent infections result in sexual transmission. In addition, infections of female mice help us determine the severity of disease caused by sexually transmitted Ebola virus, the effects on female fertility, and consequences for gestational development.

In addition to our laboratory research, we are surveying Ebola disease survivors in Sierra Leone in an effort to identify and understand the long-term health consequences, with a focus on reproductive health. Our goal is to determine if survivors have encountered post-recovery reproductive complications such as reduced fertility or problems during pregnancy.

We are also investigating whether Ebola disease survivors face greater burdens of subsequent infections by other pathogens.

Future outbreaks

The recent success of vaccination and therapeutic treatments in the current DRC Ebola disease outbreak provides hope. But a great deal about the virus remains unknown. A greater understanding of the mechanisms of how persistent infections occur will help us guide future outbreak response efforts, with the goal of reducing additional Ebola virus transmission events and providing increased care, support and safety for survivors and their communities.

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